

PCTWORLD INTELLECTUAL PROPERTY ORGANIZATION
International Bureau

INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(51) International Patent Classification ⁶ : A01N 37/18, 43/04, C07H 21/04, C12N 5/00, 5/09, 5/10, 15/09, 15/12, C12Q 1/68	A1	(11) International Publication Number: WO 99/08522 (43) International Publication Date: 25 February 1999 (25.02.99)
(21) International Application Number: PCT/US98/16816 (22) International Filing Date: 14 August 1998 (14.08.98) (30) Priority Data: 60/055,868 15 August 1997 (15.08.97) US (71) Applicants: LUDWIG INSTITUTE FOR CANCER RE-SEARCH [US/US]; 1345 Avenue of the Americas, New York, NY 10105 (US). HELSINKI UNIVERSITY LI-CENSING LTD., OY [FI/FI]; Koetilantie 3, FIN-00710 Helsinki (FI). UNIVERSITY OF GENEVA [CH/CH]; Medical Centre, Faculty of Medicine, 1, rue Michel Servet, CH-1211 Geneva 4 (CH). (72) Inventors: PEPPER, Michael, S.; 6, chemin du Chateau, CH-1245 Collonge-Bellerive (CH). ALITALO, Kari; Nyyrikintie 4A, FIN-02100 Espoo (FI). ERIKSSON, Ulf; Haegervaege 27, S-746 34 Balsta (SE). (74) Agent: EVANS, Joseph, D.; Evenson, McKeown, Edwards & Lenahan, P.L.L.C., Suite 700, 1200 G Street, N.W., Washington, DC 20005 (US).		(81) Designated States: AU, CA, CN, JP, KR, NZ, European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE). Published <i>With international search report.</i> <i>With amended claims.</i>
(54) Title: STIMULATION, MODULATION AND/OR INHIBITION OF ENDOTHELIAL PROTEOLYTIC ACTIVITY AND/OR ANGIOGENIC ACTIVITY (57) Abstract Vascular endothelial growth factor-B (VEGF-B) and vascular endothelial growth factor-C (VEGF-C) are angiogenic polypeptides. It has been shown that VEGF-B and -C are angiogenic <i>in vitro</i> especially in combination with bFGF. VEGF-C also increases plasminogen activator (PA) activity in bovine endothelial cell lines and this is accompanied by a concomitant increase in PA inhibitor-1. Addition of alpha-2-antiplasmin to bovine endothelial cells co-treated with bFGF and VEGF-C partially inhibits collagen gel invasion.		